TOPIC INFO

TOPIC:	SELECTING DISEASE-MODIFYING ANTI-RHEUMATIC DRUGS FOR RHEUMATOID ARTHRITIS
SPEAKER:	AMISH J. DAVE, MD, MPH
TITLE:	RHEUMATOLOGIST
AFFILIATION	VIRGINIA MASON MEDICAL CENTER
TIME:	30 minutes

PRACTICE GAP ANALYSIS: SELECTING DISEASE-MODIFYING ANTI-RHEUMATIC DRUGS FOR RHEUMATOID ARTHRITIS

Describe the problems or gaps in practice this activity will address:

What are you trying to change?

Currently nearly 2% of Americans have rheumatoid arthritis and appropriate diagnosis and initiation of disease-modifying antirheumatic drug within the first three months of symptom onset is now standard-of-care as per the American College of Rheumatology. Washington State is one of the worst states in the nation per the 2015 American College of Rheumatology Workforce Survey in terms of number of practicing rheumatologists to patient need. As such, in many parts of Washington State, patients can face delays of up to one year to be seen by a rheumatologist. Surrounding states, including Idaho, Wyoming, Montana, Oregon, and Alaska have similar deficits in number of rheumatologists. As such, primary care providers are managing patients with inflammatory polyarthritis, including rheumatoid arthritis, for lengthy periods of time in our region. In this talk, we aim to focus on common rheumatologic conditions (such as rheumatoid arthritis and psoriatic arthritis) seen by primary care providers and provide them with medical knowledge about diagnosis and treatment of these medical conditions.

What is the problem?

Early treatment for rheumatoid arthritis should involve patients with this autoimmune condition being diagnosed and begun on steroid-sparing disease-modifying anti-rheumatic drug therapy (DMARD) within three months of symptom onset. Primary care providers should understand what initial labs to send off for patients with inflammatory polyarthritis and how to distinguish between crystalline and non-crystalline inflammatory polyarthritis. Patients and providers should understand the incidence and prevalence (epidemiology) of rheumatoid factor (RF), anti-nuclear antibody (ANA), anti-citric citrullinated antibody (CCP), and HLA-B27 antigen studies in making a diagnosis of inflammatory polyarthritis. Primary care providers should understand the role of corticosteroids and biologic and non-biologic DMARDs in short-term and long-term management of rheumatoid arthritis and other inflammatory polyarthropathies. Primary care providers also should understand risks of infection and malignancy associated with DMARD therapy for common autoimmune conditions. Basic knowledge about risks of biologic and non-biologic DMARD therapies with pregnancy, as well as risks of corticosteroids with glycemic control in diabetes is also important.

How did you assess and/or measure these issues?

How was the educational need/practice gap for this activity identified? Place an X by each source utilized to identify the need for this activity.

Attach copies of documentation for each source indicated (REQUIRED)

* please make sure when selecting your needs assessment data and references that you highlight applicable components.

	Me	thod	Example of required document
		Previous participant evaluation data	Copy of tool and summary data
	х	Research/literature review	Abstract(s) or articles
	x	Expert Opinion	Summary
		Target audience survey	Copy of tool and summary data
		Regulatory body requirements	Requirements summary
		Data from public health sources	Abstract, articles, references
De	scribe	Other (describe) the needs of learners underlying the gaps in practice:	
		at are the causes of the gaps in practice? Check all that apply	
	х	Lack of awareness of the problem,	Poor self-efficacy,
	x	Lack of familiarity with the guideline,	x Inability to overcome the inertia of previous practice, and
		Non-agreement with the recommendations,	Presence of external barriers to perform recommendations
	Wh	y does the gap exist? Check all that apply	
	х	Lack of Knowledge competence	Lack of time to assess or counsel patients
	х	Performance-based.	Cost / Insurance/reimbursement issues
		Lack of consensus on professional guidelines	Patient Compliance Issues
		Other:	
	Wh	at do learners need to be able to know or do to be able to addr	ress the gaps in practice?
	lt ca	rheumatoid arthritis	

CME OBJECTIVES: SELECTING DISEASE-MODIFYING ANTI-RHEUMATIC DRUGS FOR RHEUMATOID ARTHRITIS

State at least three or more things that participants should be able to do after they participate in this CME activity. Please note these objectives should be measurable, specific, actionable and timely.

Upon completion of this activity, attendees should be able to:

1	Describe the roles of the rheumatologist and primary care provider in diagnosing rheumatoid arthritis
2	Identify the mechanisms of action of biologic and non-biologic disease-modifying anti- rheumatic drugs used to treat rheumatoid arthritis
3	Compare differences in outcomes and extraarticular manifestations of rheumatoid arthritis in patients with seropositive and seronegative rheumatoid arthritis
The	ACCME does not want you to use the words - think, understand, know, appreciate, learn, comprehend, be aware of, be familiar with,
etc	. as they are not measurable.
You	a can use words such as Analyze, Categorize, Classify, Compare, Conclude, Construct, Critique, Define, Demonstrate, Describe, Discuss,
Eva	luate, Identify, List, Name, Outline, Show

COMPETENCIES: SELECTING DISEASE-MODIFYING ANTI-RHEUMATIC DRUGS FOR RHEUMATOID ARTHRITIS

What ACGME or IOM related competency is associated with this activity? (check all that apply)

	х	Patient Care		Practice-Based Learning and Improvement	х	Medical/Clinical Knowledge
		Procedural Skills		Interdisciplinary Teams		Teams and Teamwork
	х	Communication Skills		Professionalism	x	Systems-based Practice
	x	Quality Improvement		Utilization of Informatics	x	Evidence-based Practice
Wh	What is the activity designed to change					
	х	Competence - (knowing how to do somet	hing)			
		Selecting this option requires the CME act	ivity	being planned provide participants with an op	port	unity to:
		hear information related to adv	ances	or best practice		
		hear examples of application in	pract	ice of information presented		
		Performance- (actually doing something)				
		Selecting this option requires the CME act	ivity	being planned provide participants with an op	port	unity to:
		• practice what they have learned	l duri	ng the CME activity		
		• receive feedback about doing w	hat tl	ney have learned during the CME activity		
		Patient Outcomes- (actually measure cha	nge ir	n patients)		
		Selecting this option requires the CME act	ivity	track change in patient outcomes:		
		• provide tangible improvements	and o	data to support overall change to patient out	ome	5
Wł	nat po	otential barriers do you anticipate attendees	s may	encounter when incorporating new objective	es int	o their practice?
	х	Lack of time to assess or counsel patients		Other – describe:		
		Cost				
		No perceived barriers				
		Lack of administrative support/resources				
		reimbursement issues				
		Insurance/				
De	scribe	e how will this educational activity address t	hese	potential barriers and the strategies used?		

RESULTS: SELECTING DISEASE-MODIFYING ANTI-RHEUMATIC DRUGS FOR RHEUMATOID ARTHRITIS please describe the results expected (outcomes) for this activity in terms of specific improvements in patient care and/or other work related to the practice of medicine. x Improvements in patient care based on evidence-based treatment Reduce Health care costs x x Streamline care of patients

MEASURING YOUR SUCCESS: SELECTING DISEASE-MODIFYING ANTI-RHEUMATIC DRUGS FOR RHEUMATOID ARTHRITIS

Will use pre-and post CME activity questionnaire to measure success.

Please provide 3 questions and answers that will asked to the audience before and after your talk. The answer to these questions should be in your presentation. Please highlight the correct answer and limit your possible answers to a maximum of 4 with only one correct answer. The others can be partially correct or wrong

Question 1. This biologic DMARD has been approved by the Food and Drug Administration for management of uveitis, hidradenitis suppurativa, rheumatoid arthritis, juvenile idiopathic arthritis, ulcerative colitis, psoriasis, and psoriatic arthritis

Ar	nswers
1	infliximab
2	etanercept
3	adalimumab
4	methotrexate
	Feedback:
	1. Infliximab: Partially Correct:
	is approved by the Food and Drug Administration for ankylosing spondylitis, pediatric and adult Crohn's disease,
	pediatric and adult ulcerative colitis, psoriatic arthritis, plaque psoriasis, and rheumatoid arthritis.
	Ankylosing spondylitis
	Pediatric and adult Crohn's disease
	Adult ulcerative colitis
	Psoriasis, and psoriatic arthritis.
	Rheumatoid arthritis,
	Reference: Ulcerative Colitis, Rheumatoid Arthritis, Ankylosing Spondylitis, Psoriatic Arthritis, Psoriasis
	2. Etanercept: Partially Correct:

		is approved by the Food and Drug Administration for rheumatoid arthritis, polyarticular juvenile idiopathic arthritis,
		psoriatic
		Arthritis, ankylosing spondylitis, and plaque psoriasis.
		Juvenile idiopathic arthritis,
		Psoriasis, and psoriatic arthritis.
		Rheumatoid arthritis,
		Reference: https://www.accessdata.fda.gov/drugsatfda_docs/nda/2003/103795s5123_EnbrelTOC.cfm
		3. Adalimumab: Correct Answer
		is the only biologic disease modifying anti-rheumatic drug (DMARD) approved by the Food and Drug Administration
		for management of all of the following conditions: of
		Hidradenitis suppurativa,
		Juvenile idiopathic arthritis,
		Psoriasis, and psoriatic arthritis.
		Rheumatoid arthritis,
		Ulcerative colitis,
		Uveitis,
		Reference: https://www.accessdata.fda.gov/drugsatfda_docs/label/2018/125057s410lbl.pdf
		4. Methotrexate: Wrong Answer.
		has been approved for various medical conditions including rheumatoid arthritis and acute lymphoblastic
		leukemia, but not for uveitis or hidradenitis suppurativa. It has long been used for management of inflammatory
		bowel diseases and psoriasis as well as psoriatic arthritis.
		Reference: https://www.europeanpharmaceuticalreview.com/news/107390/fda-approves-new-line-of-
		methotrexate-products-for-rheumatoid-arthritis/
Qu	estior	2: Which biologic DMARD is PEGylated and is the choice for patients with rheumatoid arthritis who are pregnant or breastfeeding?
	Ans	wers
	1	<mark>certolizumab</mark>
	2	secukinumab
	3	ixekizumab
	4	apremilast
	5	adalimumab
		Feedback: Please provide a detail feedback (MOC) requirements for above questions.
		1. Certolizumab: correct Answer
		is a PEGylated molecule with a large polyethylene glycol moiety in its chemical structure that limits its ability to
		cross the placenta or into breast milk. Per the Mother-to-Baby educational resource based at the University of
		California San Diego, animal studies found no difference in fertility when taking certolizumab. There are no formal
		studies looking at risks to pregnancy with any biologic medication in rheumatoid arthritis. However, minimal
		certolizumab has been detected in children delivered to women on this biologic DMARD medication during their
		gestation.
		2. Secukinumab: Wrong Answer.

are not FDA-approved medications for rheumatoid arthritis. Adalimumab crosses the placenta in considerable amounts and less preferred as compared to certolizumab during pregnancy

3. Ixekizumab: Wrong Answer.

are not FDA-approved medications for rheumatoid arthritis. Adalimumab crosses the placenta in considerable amounts and less preferred as compared to certolizumab during pregnancy

4. Apremilast: Wrong Answer.

are not FDA-approved medications for rheumatoid arthritis. Adalimumab crosses the placenta in considerable amounts and less preferred as compared to certolizumab during pregnancy

5. Adalimumab: Wrong Answer.

References:

• Mariette X, et al. 2017. Lack of placental transfer of certolizumab pegol during pregnancy: results from CRIB, a prospective, post marketing, pharmacokinetic study., Ann Rheum Dis; 0:1–6. doi:10.1136/annrheumdis-2017-212196

• Clowse, ME, et al. 2018. Pregnancy Outcomes after Exposure to Certolizumab Pegol: Updated Results from a Pharmacovigilance Safety Database. Arthritis Rheumatol. Accepted Author Manuscript. doi:10.1002/art.40508

• Clowse MEB, et al. 2017. Minimal to no transfer of certolizumab pegol into breast milk: results from CRADLE, a prospective, post marketing, multicenter, pharmacokinetic study. Ann Rheum Dis; 76:1890–1896.

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https://www.cimzia.com/sites/default/files/docs/CIMZIA_Current_COL_03-2018_effective_21_March_2018.pdf Accessed 16 April 2018.

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• Habal FM and Huang VW. 2012. Review article: a decision-making algorithm for the management of pregnancy in the inflammatory bowel disease patient. Aliment Pharmacol Ther 35(5):501-15.

• Khan N, et al. 2014. Safety of anti-TNF therapy in inflammatory bowel disease during pregnancy. Expert Opin Drug Saf; 13(12):1699-708.

 Mahadevan U, et al. 2011. The London Position Statement of the World Congress of Gastroenterology on Biological Therapy for IBD with the European Crohn's and Colitis Organization: pregnancy and pediatrics. Am J Gastroenterol. 106(2):214-23

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• Wakefield I, et al. 2011. The use of surrogate antibodies to evaluate the developmental and reproductive toxicity potential of an anti-TNF alpha PEGylated Fab' monoclonal antibody. Toxicol Sci 122(1):170-6.

• Weber-Schoendorfer C; network of French pharmacovigilance centers, et al. 2015. Pregnancy outcome after TNF- α inhibitor therapy during the first trimester: a prospective multicenter cohort study. Br J Clin Pharmacol; 80(4):727-39.

• Wolf D and Mahadevan U. 2010. Certolizumab use in pregnancy: low levels detected in cord blood. Arthritis Rheum [abstract] 62Suppl10:718.

Question 3: Patients with seropositive rheumatoid arthritis are less likely to develop mononeuritis multiplex, corneal melt, rheumatoid lung disease, and Felty's syndrome compared to patients with seronegative rheumatoid arthritis. True or False?

1	True
2	False
	Feedback: Please provide a detail feedback (MOC) requirements for above questions.
	False. <mark>(correct Answer</mark>)
	Patients with seropositive rheumatoid arthritis involving either a positive rheumatoid factor and/or positive citric
	citrullinated antibody are more likely to have aggressive complications of rheumatoid arthritis, including risks of rheumatoid
	vasculitis, Mononeuritis multiplex, corneal melt, uveitis and iritis, rheumatoid lung disease (including either non-specific
	interstitial pneumonitis or usual interstitial pneumonia) or pulmonary nodulosis (Caplan's disease). Felty's syndrome – a tria
	of neutropenia, splenomegaly, and long-standing rheumatoid arthritis – is almost always seen in patients with rheumatoid
	factor-positive rheumatoid arthritis for many years.
	Reference:
	Nat Rev Rheumatol. 2015 Jan;11(1):8-9. doi: 10.1038/nrrheum.2014.194. Epub 2014 Nov 18.
	Best Pract Res Clin Rheumatol. 2004 Oct;18(5):631-45. Review .